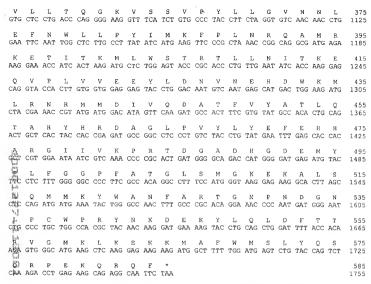
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$ \begin{array}{cccccccccccccccccccccccccccccccccccc$	15 3 45
P T A G A G W S M R W I L C W S L T L CCC ACA GGA GGA GGC TGG AGC ATG AGG TGG ATT CTG TGC TGG AGC CTC ACC CTC TGC AGC TGC TGC TGC TGC TGC TGC TGC TGC TGC T	
. L M A Q T A L G A L H T K R P Q V V T T CTG ATG GCG CAG ACG GCC TTG GGT GCC TTG CAC ACC AAG AGG CCT CAA GTG GTC ACC AAG	
Y G T L Q G K Q M H V G K T P I Q V F I TAT GGA ACC CTG CAA GGA AAA CAG ATG CAT GTG GGG AAG ACA CCC ATC CAA GTC TTT T	
G V P F S R P P L G I L R F A P P E P B B B B G C C C T T C T C AGA C C T C T AGG T AT C C T AGG T T G C A C C T C T C AGA C C C C T C T C AGA C C C C C C C C C C C C C C C C C	95 CG 285
P W K G I R D A T T Y P P G C L Q E S  AG CCC TGG AAA GGA ATC AGA GAT GCT ACC ACC TAC CCG CCT GGG TGC CTG CAG GAG TC  LL  LL	
G GC CAG CTG GCC TCG ATG TAC GTC AGC AGG CGG GAA CGG TAC AAG TGG CTG CGC TTG	
AGC GAG GAC TGT CTG TAC CTG AAC GTG TAC GCG CCG GCG CGC GCG CCC GGG GAT CCC CA	
$\stackrel{L}{\mid A \mid}$ P V M V W F P G G A F I V G A A S S ) QWG CCA GTG ATG GTC TGG TTC CCG GGA GGC GCC TTC ATC GTG GGC GCT GCT TCT TCG T/L C)	
G S D L A A R E K V V L V F L Q H R I GAG GGC TCT GAC TTG GCC CGC GAG AAA GTG GTG CTG GTG TTT CTG CAG CAC AGG CT	
GGC ATC TTC GGC TTC CTG AGC ACG GAC GAC GAC GCG CGC GGG AAC TGG GGG CTG CTG AGC ACG GAC GAC GCG CGC GGG AAC TGG GGG CTG CTG CTG GGG CTG CTG CTG CT	
D Q M A A L R W V Q E N I A A F G G G G GA GAC CAG ATG GCG GTG GTG GTG CAG GAG AAC ATC GCA GCC TTC GGG GGA GAC CC	
G N V T L F G Q S A G A M S I S G L M $^{\rm h}$ GGA AAT GTG ACC CTG TTC GGC CAG TCG GGG GCC ATG AGC ATC TCA GGA CTG ATG ATG	
S P L A S G L F H R A I S Q S G T A L F TCA CCC CTA GCC TCG GCT CTC CAT CGG GCC ATT TCC CAG AGT GGC ACC GCG TTA TCC CAG AGT GCC ACC GCG TTA TCC CAG ACC GCG TTA TCC CAG ACC GCG TTA TCC CAG ACC ACC GCG TTA TCC CAG ACC GCG TTA TCC CAG ACC ACC GCG TTA TCC CAG ACC ACC GCG TTA TCC CAG ACC ACC ACC ACC ACC ACC ACC ACC A	
R L F I T S N P L K V A K K V A H L A GAGA CTT TTC ATC ACT AGA CCA CTG AAA GTG GCC AAG AAG GTT GCC CAC CTG GCT GC	
C N H N S T Q I L V N C L R A L S G T F G C AAC CAC AAC AGC ACC CG ACC CTG GTA AAC TGC CTG AGG GCA CTA TCA GGG ACC AAC CAC AAC AGC ACC CTG CTG AGG GCA CTA TCA GGG ACC AAC CAC ACC CAC CAC CAC CAC C	
V M R V S N K M R F L Q L N F Q R D P GTG ATG CGT GTG TCC AAC AAG ATG AGA TTC CTC CAA CTG AAC TTC CAG AGA GAC CCG GA	
E I I W S M S P V V D G V V I P D D P I GAG ATT ATC TGG TCC ATG AGC CCT GTG GTG GAT GGT GTG GTG ATC CCA GAT GAC CCT TTG GAG GAT GAC CCT TTG GAC GAT GAC CCT TTG GAG GAT GAC CCT TTG GAC GAT GAC CCT TTG GAG GAT GAC CCT TTG GAC GAC GAT GAC CCT TTG GAC GAC GAC GAC GAC GAC GAC GAC GAC GA	



GGGTGCTATGCAGGAAGGAGCCAAAGAGGGGTTTGCCCCCACCATCCAGGCCCTGGGGAGACTAGCCATGGACATACC

TGGGGACAAGAGTTCTACCCAAGGGCGAATTCGTTTAAACCTGCAGGACTAG

Figure 1B

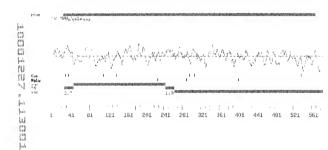


Figure 2

## Protein Family / Domain Matches, HMMer version 2

```
Searching for complete domains in PFAM
    hmmpfam - search a single seg against HMM database
    HMMER 2.1.1 (Dec 1998)
    Copyright (C) 1992-1998 Washington University School of Medicine
    HMMER is freely distributed under the GNU General Public License (GPL).
    HMM file:
Sequence file:
     IMM file: /prod/ddm/seqanal/PFAM/pfam5.5/Pfam
sequence file: /prod/ddm/wspace/orfanal/oa-script.26660.seq
     Query: 18903
    Scores for sequence family classification (score includes all domains):
  Model
          Description
                                                       Score
                                                              E-value N
  COesterase Carboxylesterases
                                                        558.6 4.1e-164 1
Parsed for domains:
Model Domain seq-f seq-t hmm-f hmm-t
                                                 score E-value
 COesterase 1/1
                   25 569 ...
                                    1 612 () 558.6 4.1e-164
Alignments of top-scoring domains:
COesterase: domain 1 of 1, from 25 to 569: score 558.6, E = 4.1e-164
111
                    *->mvllllfLllllllliavlaaakaspedpllVatnnVlcGkvrGvnek
                        +1+ +L 1 1 ++ +1+a+++ ++ V t++ G++ G++
4, 1
        18903
                25
                      RWILCWSLTLCLMAOTALGALHT---KRPQVVTKY---GTLOGKOMH 65
ā
Lak
                   tdngeqsvysFlGIPYAePPVGnLRFkaPqPYkepWsdvldAtkyppsCl
100
                        ++ +FlG+P+ PP+G LRF +P+P +epW++++dAt+ypp Cl
        18903
                66 VGKT--PIQVFLGVPFSRPPLGILRFAPPEP-PEPWKGIRDATTYPPGCL 112
L
                   QdddfgfslsdLKvalkmlslgwnklvg....lklsEDCLYLNVvtPknt
13
                   Q + +g+ 1 +++ + + 1 +sedClylnvy P+ +
        18903
               113 Q-ESWGQ-----LASMYVSTRETVkwLRFSEDCLYLNVYAPARA 150
                   kpnsklPVmVwIhGGGFmfGsghslplslYdgeslaregnVIvVsiNYRL
                   + + +1PVmVw +GG+F +G++ s+Y g+ la++++V++V ++ RL
        18903
               151 PGDPQLPVMVWFPGGAFIVGAA----SSYEGSDLAAREKVVLVFLQHRL 195
                   GplGFLstgddklpgsGNyGL1DQrlALkWVqdNIaaFGGDPnsVTifGe
                   G++GFLst+d+++ GN+GL1DQ +AL+WVq+NIaaFGGDP++VT+fG+
        18903
               196 GIFGFLSTDDSHAR--GNWGLLDQMAALRWVQENIAAFGGDPGNVTLFGQ 243
                   SAGaaSVsllllsngGDNppsskgLFhRAIsqSGsalspwaiqsesnarg
                   SAGa+S+s 1++s p++ gLFhRAIsqSG+al +i+s+
        18903
               244 SAGAMSISGLMMS----PLA-SGLFHRAISQSGTALFRLFITSNP--LK 285
                   rakelarl1GCnetsssel1dCLRsksaeeLleatrsfllfeyvpflplf
                    ak++a 1+GCn++s 1+ CLR s + + + + + f + +f
               286 VAKKVAHLAGCNHNSTQILVNCLRALSGTKVMRVSNK-MRFLQLNFQRDP 334
        18903
                   1....aFgPvvDGdDapeafipedPeelikeGkfadvPyliGvtkdEGqy
                   ++ + +PvvDG+ +ip+dP+ 1+ +Gk + vPyl Gv++ E+ +
       18903
               335 EeiiwSMSPVVDGV----VIPDDPLVLLTQGKVSSVPYLLGVNNLEFNW 379
```

18	903 380	faamllnasskgedelkketnpdvwlellkyllfyasealnikdMddlad ++++ +++ + ket ++ 1+ 11+ ++ + LLPYIMKFPLNR-QAMRKETITK-MLWSTRTLLN-ITKEQ	18
18	903 419	kvlekYpgdvddfsvesrkpnlqdmltDllFkcptrvaadlhakhggsPv v e+Y++ v +++++++++ D+F++++++++++++++++++++++	67
18	903 468	YaYvfdhpasfgigQflakrvdpefggavHgdEiffvFgnpllkeqlyka Y+Y+f h+a+ + v+p+++ga+HgdE++f+Fg p+ ++ 1 YLYEFEHHARGIIVKPRTDGADHGDEMYFLFGGPFATG-LS 5	107
18	903 508	teeeeksssktmmnywanFAktGnPnngtsnglvvWpkytseeqkYsll1 e+++s +mm+ywanFA+tGnP n++ +1+ Wp y+++e +1+ MGKEKALS-LQMMKYWANFARTGNP-NDGNLPCWPRYNKDEKYLQL 5	51
18	903 552	llttitagklkardprkvlcnfw<-* +tt +klk+++ ++fw DFTTRVGMKLKEKKMAFW 569	
// Searching	for comp	lete domains in SMART	

```
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    Fbh18903FL - Import - vector trimmed
    to: FrGcgManager_76 JOA3WXZ1 check: 1319 from: 1 to: 2456
    Z34105 in Patent Nucleotide
    Symbol comparison table: /ddm_local/gcg/gcg_9.1/gcgcore/data/rundata/nwsgapdna.cmp
    CompCheck: 8760
        Gap Weight: 12 Average Match: 10.000
Length Weight: 4 Average Mismatch: 0.000
             Quality: 13796
                                      Length: 2746
               Ratio: 6.957
                                       Gaps: 14
    Percent Similarity: 92.617 Percent Identity: 92.617
          Match display thresholds for the alignment(s):
                    | = IDENTITY
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234105 101 CATTTCGCCTTGCTGACGGCGTCGAGGCCCTGGCCAGACATGTCCACAGGG 150
         4 TTAGCCAATTCGGCCGAGGCCTCCCGCCCCAGTA.CTTGCTGGCAGGGAT 52
           151 TTCTCCTTCGGGTCCGGGACTCTGGGCTCCACCACCGTGGCCGCCGGCGG 200
        53 TAAGAGCAGA.TAAAAGTGTGCTCACACACTGTAGACACGGCTACCATGC 101
            1 1111 1 11 111 1 1 131 111 11
        201 GACCAGCACAGGCGGCGTTTTCTCCTTCGGAACGGGAACGTCTAGCAACC 250
        102 CATC.....CACAGTGTTGCCATCCACAGTGTTGCCATCACTCCTGC. 143
           251 CTTCTGTGGGGCTCAATTTTGGAAATCTTGGAAGTACTTCAACTCCAGCA 300
        144 .CCACAGCAGGAGCT..GGCTGGAGCATGAGGTGGATTCTGTG..CTGGA 188
            301 ACTACATCTGCTCCTTCAAGTGGTTTTGGAACCGGGCTCTTTGGATCTAA 350
```

Figure 4A

189	GCCTCACCCTCTGCCTGATGGCGCAGACGGCCTTGGGTGCCTTGCAC	235
351	ACCTGCCACTGGGTTCACTCTAGGAGGA . ACAAATACAGGTGCCTTGCAC	399
236		285
400	ACCAAGAGGCCTCAAGTGGTCACCAAATATGGAACCCTGCAAGGAAAACA	449
286	GATGCATGTGGGGAAGACACCCATCCAAGTCTTTTTAGGAGTCCCCTTCT	335
450		499
336		385
500		549
	CCCTGGAAAGGAATCAGAGATGCTACCACCTACCCGCCTG	425
	CCCTGGAAAGGAATCAGAGATGCTACCACCTACCCGCCTGGATGGA	-
550		299
	:	
426		465
700	TGGGGCTACAGGTGCCTGCAGGAGTCCTGGGGCCAGCTGGCCTCGATGTA	749
466		515
750	CGTCAGCACGCGGAACGGTACAAGTGGCTGCGCTTCAGCGAGGACTGTC	799
516		565
800	TGTACCTGAACGTGTACGCGCGGGCGCGCGCGCGCGGGATCCCCAGCTG	849
566		615
	тининининининий интипи	899
	TTCGTACGAGGCTCTGACTTGGCCGCCCGGAGAAAGTGGTGCTGGTGT	949
	TTCTGCAGGACAGGCTCGGCATCTTCGGCTTCCTGAGCACGGACGACAGC	
		999
	CACGCGCGCGGGAACTGGGGGCTGCTGGACCAGATGGCGGCTCTGCGCTG	

Figure 4B

1050	GGTGCAGGAGAACATCGCAGCCTTCGGGGGAGACCCAGGAAATGTGACCC	1099
816	TGTTCGGCCAGTCGGCGGGGCCATGAGCATCTCAGGACTGATGATGTCA	865
1100	######################################	1149
866	CCCCTAGCCTCGGGTCTCTTCCATCGGGCCATTTCCCAGAGTGGCACCGC	915
1150	CCCCTAGCCTCGGGTCTCTTCCATCGGGCCATTTCCCAGAGTGGCACCGC	1199
916	GTTATTCAGACTTTTCATCACTAGTAACCCACTGAAAGTGGCCAAGAAGG	965
	$\begin{tabular}{lllllllllllllllllllllllllllllllllll$	
	TTGCCCACCTGGCTGGATGCAACACACACACACACACACA	
	TTGCCCACCTGGCTGGATGCAACCACAACAGCACACAGATCCTGGTAAAC	
	TGCTGAGGCACTATCAGGGACCAAGGTGATGCGTGTGTCCAACAAGAT	
1066	. $ \\ \textbf{GAGATTCCTCCAACTGAACTTCCAGAGAGACCCGGAAGAGATTATCTGGT} $	1119
1350		1399
1116	CCATGAGCCCTGTGGTGGTGTGGTGTTCCCAGATGACCCTTTGGTG	1165
1400	CCATGAGCCCTGTGGTGGATGGTGTGGTGATCCCAGATGACCCTTTGGTG	1449
	CTCCTGACCCAGGGGAAGGTTTCATCTGTGCCCTACCTTCTAGGTGTCAA	
	CTCCTGACCCAGGGGAAGGTTTCATCTGTGCCCTACCTTCTAGGTGTCAA	
	CAACCTGGAATTCAATTGGCTCTTGCCTTATATCATGAAGTTCCCGCTAA	
1500		1550
	CGCACCCTGTTGAATATCACCAAGGAGCAGGTACCACTTGTGGTGGAGGA	
	GTACCTGGACAATGTCAATGAGCATGACTGGAAGATGCTACGAAACCGTA	
	Figure 4C	1018

766 GGTGCAGGAGAACATCGCAGCCTTCGGGGGAGACCCAGGAAATGTGACCC 815

rigure 4C

1416	TGATGGACATAGTTCAAGATGCCACTTTCGTGTATGCCACACTGCAGACT 146	5
1619	TGATGGACATAGTTCAAGATGCCACTTTCGTGTATGCCACACTGCAGACT 166	8
1466		5
1669		3
	:	
	· · · · · · · · · · · · · · · · · · ·	
	CAACTTTGCCCGCACAGGAAACCCCAATGATGGGAATCTGCCCTGCTGGC 171	
1684		5
1716		5
1717	CACGCTACAACAAGGATGAAAAGTACCTGCAGCTGGATTTTACCACAAGA 1766	5
1766		5
1767		5
	CCAGTCTCAAAGACCTGAGAAGCAGAGGCAATTCTAAGGGTGGCTATGCA 1865	
	GGAAGGACCAAAGAGGGGTTTGCCCCCACCATCCAGGCCCTGGGGAGAC 1915	
	GGAAGGAGCCAAAGAGGGGTTTGCCCCCACCATCCAGGCCCTGGGGAGAC 1916	
	TAGCCATGGACATACCTGGGGACAAGAGTTCTACCCAAGGGCGAATTCGT 1965	
	TAGCCATGGACATACCTGGGGACAAGAGTTCTACCCACCCCAGT 1960	
1966	TTAAACCTGCAGGA.CTAG	
061	TITLE CAR CONCOLOGA COMO COMO COMO COMO CARCA LA COMA CARCA	

Figure 4D

```
ALIGN calculates a global alignment of two sequences
 version 2.0uPlease cite: Myers and Miller, CABIOS (1989)
                                  COE-2
                                 PRO873
 scoring matrix: BLOSUM50, gap penalties: -12/-2
 62.4% identity;
                 Global alignment score: 2271
COE-2 M------PST-----VLPSTVLPSLLPTAG
                                       PROST3 MSTGFSFGSGTLGSTTVAAGGTSTGGVFSFGTGTSSNPSVGLNFGNLGSTSTPATTSAPS
     20
            3.0
                   40
                          50
                                  60
 COE-2 AGWSMRWILCWSLTLCLMAQTALGALHTKRPQVVTKYGTLQGKQMHVGKTPIOVFLGVPF
      ....
PROWAS SGFGTGLFGSKPATGFTLGGTNTGALHTKRPQVVTKYGTLQGKQMHVGKTPIQVFLGVPF
            70
                  80
                          90
                                100
                                        110
            90
                   1.00
                          110
 COF-7 SRPPLGILRFAPPEPPEPWKGIRDATTYPPG-----
      PROXIX SRPPLGILRFAPPEPPEPWKGIRDATTYPPGWSLALSPGWSAVARSRLTATSASRVOASL
           130
                  140
                         150
                                160
                                        170
                   120
                          130
                                 140
 COE-2 -----CLQESWGQLASMYVSTRERYKWLRFSEDCLYLNVYAPARAPGDPOLPVM
              PROT13 LPQPLSVWGYRCLQESWGQLASMYVSTRERYKWLRFSEDCLYLNVYAPARAPGDPQLPVM
                  180
                          190
                                  200
 COE-2 VWFPGGAFIVGAASSYEGSDLAAREKVVLVFLQHRLGIFGFLSTDDSHARGNWGLLDOMA
      PLOSTS VWFPGGAFIVGAASSYEGSDLAAREKVVLVFLQHRLGIFGFLSTDDSHARGNWGLLDQMA
           250
                       270
                              280
                                       290
    220
                          250
            230
                   240
                                  260
                                          270
 COF-Z ALRWVQENIAAFGGDPGNVTLFGQSAGAMSISGLMMSPLASGLFHRAISQSGTALFRLFI
      PLOYS ALRWYQENIAAFGGDPGNYTLFGQSAGAMSISGLMMSPLASGLFHRAISQSGTALFRLFI
           310
                  320
                         330
                                 340
                                        350
    280
            290
                   300
                          310
                                  320
 CCE-2 TSNPLKVAKKVAHLAGCNHNSTOILVNCLRALSGTKVMRVSNKMRFLOLNFORDPEEIIW
      PROST3 TSNPLKVAKKVAHLAGCNHNSTQILVNCLRALSGTKVMRVSNKMRFLQLNFQRDPEEIIW
           370
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                         390
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Figure 5A

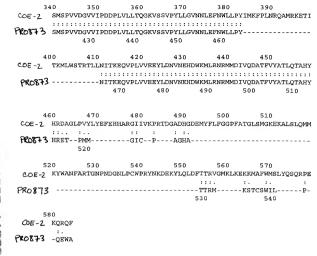
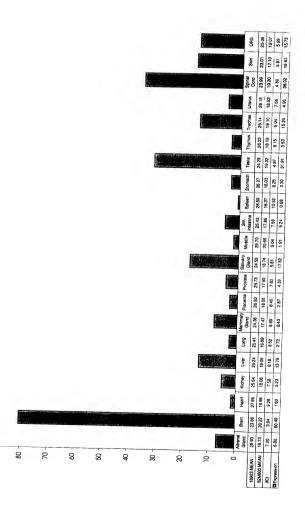


Figure 5B

8



									Hu Lund		25.805	15.87	9 835
									Hu Liver	47.50	25.43	18.22	12,
								SECOND	Hu Kidney	20 455	20 100	0.35	25.0
									Hu. Heart	28.28	17 055	10 405	
							100		Hu Spinal	23.78	18.205	5 485	24 045 4004
									Hu Brain	21.27	19 245	2 025	377 610146
									MK liver	27.39	18 22	9.17	2 FRBOAT20
									muscle	37.235	18.91	18 325	0.0046802
									LV ear	37 44	17 655	19 785	0 00170122
								MC bose	skin	36 89	18.295	18 595	0 00388138
									MK kidney	33 735	17.78	15 955	0 02419385
								MK spinal MK sciatio	nerve	9	17.37	22.63	0.00023677
								MK spinal	cord	35.89	18 71	17.18	0.01035007
								3	פונס עוני	9	17 375	22 625	0.00023759
_									Valor Valor	36.935	50.366	16.57	600000000
2	350	300	250 -	500	150	100	- 09	0		Ideuts Average Ct	TIN Average Ct	Gelfa Ci	Treature Extression   0.015/9689  0.0023/39  0.000267  0.0002677  0.02419385  0.00388138  0.00170122  0.0048802  2.66604738  2.72 5.146745  0.0048802  2.146745  0.0048802

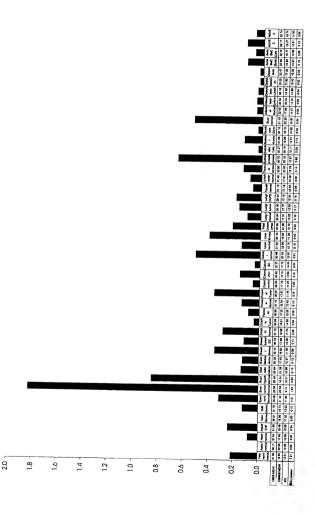


Figure 8B

Tissue		M803 MEAN •Ct	Ex	pression
Fetal heart/normal	31.89	19.68	12.21	0.21
Aorta / normal	36.11	22.53	13.58	0.00
Heart normal	31.54	17.89	13.65	0.08
Heart/ CHF	31.50	19.45	12.05	0.24
Vein/ Normal	36.01	18.99	17.02	0.00
SMC (Aortic)	31.76	18.74	13.02	0.12
Spinal cord/Normal	33.06	21.40	11.66	0.31
Brain cortex/ Normal	29.68	20.58	9.10	1.83
Brain hypothalamus/ Normal	29.45	19.24	10.21	0.85
Glial cells (Astrocytes)	33.94	21.09	12.86	0.13
Brain/"Glioblastoma Breast" Normal	30.39	17.43	12.97	0.13
	30.10	18.60	11.50	0.35
Breast tumor/IDC OVARY Normal	30.75	17.66	13.09	0.11
OVARY/ Normal OVARY/ Tumor	31.73	19.95	11.79	0.28
Pancreas	33.59	18.91	14.68	0.04
Prostate/ Normal	30.88	17.33	13.55	0.08
Prostate/Tumor	31.16	18.34	12.83	0.14
Colon≢ normal	29.03	17.57	11.46	0.35
Colon/furnor	32.03	17.78	14.26	0.05
Colon#BD	30.80	18.15	12.66	0.16
Kidne Winormal	32.37	17.72	14.65	0.04
Liver/ ggrmal	30.98	20.03	10.95	0.51
Liver fibrosis	31.82 30.10	19.08	12.75	0.15
Fetal Liver/normal	33.55	18.82	11.29	0.40
Lung / normal	30.54	21.39 17.31	12.16	0.22
Lung/ tumor	30.30	17.31	13.23	0.10
Lung/ COPD	29.44	17.77	12.53	0.17
Tonsil/ normal	31.13	17.10	12.34 13.94	0.19
Lymph node/ normal	31.40	17.19	13.49	0.06
Thymus/ normal	32.80	20.03	12.78	0.09 0.14
Epithelial Cells (prostate)	30.70	20.13	10.57	0.14
Endothelial Cells (aortic)	35.27	20.16	15.11	0.03
Skeletal Muscle/ Normal	31.06	18.26	12.81	0.03
Fibroblasts (Dermal)	33.10	18.18	14.92	0.14
Skin/ normal	31.16	20.29	10.87	0.54
Adipose/ Normal	32.43	17.96	14.47	0.04
Osteoblasts (primary)	34.84	20.44	14.40	0.05
Osteoblasts (Undiff)	33.19	18.50	14.69	0.04
Osteoblasts(Diff)	33.23	17.83	15.40	0.02
Osteoclasts	32.47	17.25	15.22	0.02
Aorta SMC (Early)	32.50	19.53	12.97	0.03
Aorta SMC (Late )	32.72	18.77	13.96	0.06
HUVEC	32.17	19.27	12.91	0.13
HMVEC	32.74	18.74	14.00	0.06

40.00

40.00

0.00